Native coding sequ Human alpha-1 anti Thyroid cancer rel Human cDNA differe

Thyroid cancer rel Sequence encoding Sequence encoding Human alphal-anti-

> AAN91077 AAN40078

Nucleotide sequenc Human alphal-antit CDNA encoding nove Nucleotide sequenc Alpha-1-antitrypsi Human alpha-1-antitrypsi

AAX83548 AAZ45928 AAZ45933 AAZ45935 AAZ45936 AAZ45936 AAZ45936 AAZ45936 AAZ45936 AAZ45936 AAZ45936 AAZ45936 AAZ45937 AAN97217

6981 7054 7405 11189 1352 Sequence encoding Sequence of human Sequence encoding Osteoarthritis tis

AAQ23746 AAQ03184 AAN50425 AAN50540 AAN50021 AAH23089

1396

429 4228.6 4227.6 427.7 425.8 425.8 424.2 424.2 424.2 421.6

Sequence of fusion

Protease inhibitor Recombinant squirr Alpha-1-antitrypsi

ABL59152 AAT79493 AAT78180

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AAN91078

Alpha-l antitrypsi Entire sequence of

Sequence of alpha-Sequence of alpha-

Human alpha 1-anti

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DNA sequence encod
DNA encoding human
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Copyright (c) 1993 - 2002
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malaria; emphysema; asthma; chronic obstructive pulmonary disease; cystic fibrosis; otitis media; otitis external; HIV; psoriasis; eczema; human immunodeficiency virus; atopic dermatitis; muscular dystrophy; herpes; ulceration; sepsis; rheumatoid arthritis; periodontal disease; tumour metastasis; osteoporosis; Paget's disease; scleroderma; glomerulonephritis; hypertension.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         rSLAP1; gene; ds; Alzheimer's disease; tumour angiogenesis;
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/standard_name= "Ribosome binding site"
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ALIGNMENTS
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          This invention relates to a novel fusion protein comprising a first protease inhibitor comprising an alphal-antitrypsin or its functionally active portion and a second protease inhibitor or its functionally active protein. The fusion proteins of the invention may act as an inhibitor of protease activity. The fusion protein of the invention is useful for inhibiting protease activity associated with a disorder such as emphysema, asthma, chronic obstructive pulmonary disease, cystic fibrosis, othis media, othis external or HIV infection, or for treating an individual suffering from or at risk for a disease or
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Novel fusion protein useful for inhibiting protease activity associated with a disorder such as emphysema, asthma, comprises a first protease inhibitor comprising alpha 1-antitrypsin and a second protease inhibitor
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       disorder involving unwanted protease activity. The proteins are useful for treating dermatological diseases such as atopic dermatitis, eczema
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tive 0; Mismatches
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/note= "SLPI coding region"
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/*tag= d
/note= "linking codon"
1197..1517
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Matches 1525; Conservative
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                                                        301 AAGAGTTGTTGAGAACTTTGAATCAACCTGATTCTCAATTGCAATTAACTACTGGTAACG
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                                                                                                     TATATCATAGTGAGGCTTTTACCGTTAATTTTGGTGATACTGAGGAAGCTAAAAAAGCAAA
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rN-TAP1; gene; ds; Alzheimer's disease; tumour anglogenesis; malaria; emphysema; asthma; chronic obstructive pulmonary disease; cystic fibrosis; ottitis media; ottitis external; HIV; psoriasis; eczema; human immunodeficiency virus; atopic dermatitis; muscular dystrophy; herpes; ulceration; sepsis; rheumatoid arthritis; periodontal disease; tumour metastasis; osteoporosis; Paget's disease; scleroderma;
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             1381 GTAAGTGTCCAGTTACTTACGGTCAATGTTTGATGTTGAACCCAACCTTCTGTGAAA
                                               TGGACGGTCAATGTAAGAGAGTCTTGAAGTGTTGTATGGGTATGTGTGGTAAGTCCTGTG
                                                                                                                                                                                                                                                                                                                                                                                                                           /standard_name= "Ribosome binding site"
9.1577
                                                                                                                                                                                                                                                                                                                                                                                                                                                               /product- "rTAP1 fusion protein"
                                                                                                                                                                                                                                          DNA sequence encoding rN-TAP1 fusion protein.
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/note= "TIMP-1 coding region"
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/note= "AAT coding region"
1194..1196
                                                                                                      TTTCCCCAGTCAAGGCCTAGTCGAC 1525
                                                                                                                                                                                                                                                                                                                                                                                               Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Pemberton P;
                                                                                                                                                                                                                                                                                                                                      glomerulonephritis; hypertension
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                /note= "linking
                                                                                                                                                                        ABK88027 standard; DNA; 1582
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/*tag= e
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                                                                                                                                                                                                                    (first entry)
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         /*tag=
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              WPI; 2002-500631/53
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                                                                                                                                                                                                                                                                                                                                                            Homo sapiens.
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                                                                                                                                                                                                                                                                                                                                                                        Synthetic
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                                                                                                                                                                                                ABK88027;
                      1381
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                                                                   1441
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This invention relates to a novel fusion protein comprising a first protease inhibitor comprising an alphal-antitrypsin or its functionally active portion and a second protease inhibitor or its functionally active portein. The fusion proteins of the invention may act as an inhibitor of protease activity. The fusion protein of the invention is useful for inhibiting protease activity associated with a disorder such as emphysema, asthma, chronic obstructive pulmonary disease, csuch as emphysema, asthma, chronic obstructive pulmonary disease, csuch as emphysema, asthma, chronic obstructive pulmonary disease, csuch as emphysema, asthma, chronic obstructive pulmonary disease or for treating an individual suffering from or at risk for a disease or discate involving unwanted protease activity. The proteins are useful for treating dermatological diseases such as atopic dermatitia, eczema and psoriasis, in infilammatory responses to viral infection, and for treating herpes infection, corneal or epidermal ulceration, chronic tumour metastasis and tumour anglogenesis, gastric ulceration, corneal or condemnation and disease, tumour metastasis and tumour anglogenesis, gastric ulceration,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ö
                                                                                                                                                                                                                                                                                                                                       osteoporosis, Paget's disease, glomerulonephritis, scleroderma, malaria, bacterial infection, Alzhalmer's disease, hypertension and muscular dystrophy. The present sequence represents the DNA encoding the rN-TAPI fusion protein of the invention.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      999
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  241 TAGAAGGITTAAACTITAATTIGACCGAAATCCCAGAAGCCCAAATTCACGAGGGTTTIC 300
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               361 GTTTATTTTTGTCTGAAGGTTTAAAATTGGTTGACAAATTCCTAGAAGACGTCAAGAAAC 420
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       TATATCATAGTGAGGCTTTTACCGTTAATTTTGGTGATACTGAGGAAGCTAAAAAGCAAA 480
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             661 AAGTICCAATGATGAAAAGACTGGGTATGTTCAATATTCAACATTGCAAAAATTAAGTT 720
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      501 CTTTCGAGGTTAAAGATACTGAAGAGGAAGATTTTCATGTTGATCAAGTTACTACTGTCA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Score 1197.4; DB 24; Length 1582;
Pred. No. 5.5e-291;
0; Mismatches 1; Indels 0;
                                                                                                                                                                                                                                                                                                                                                                                                                                           Sequence 1582 BP; 464 A; 334 C; 329 G; 455 T; 0 other;
Example 3; Page 95-96; 134pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               78.5%;
99.9%;
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rTAP1; gene; ds; Alzheimer's disease; tumour angiogenesis; malaria; emphysema; asthma; chronic obstructive pulmonary disease; cystic fibrosis; otitis media; otitis external; HTV; psorlaais; eczema; human immunodeficiency virus; atopic dermatitis; muscular dystrophy; herpes; ulceration; sepsis; rheumatoid arthritis; periodontal disease; tumour metastasis; osteoporosis; Paget's disease; scleroderma; glomerulonephritis; hypertension.
                                                                                                                                                                                                                 ATTTGAGTGGTGTTACTGAAGAAGCTCCATTAAAATTGAGTAAAGCTGTTCACAAAGCCG 1020
                                                                                                                                                                                                                               TCTTAACTATTGATGAAAAGGGTACCGAGGCCGCCGCGCGTATGTTCCTGGAAGCTATTC 1080
                                                                                                                                                                                                                                                                        CAATGAGCATTCCACCAGAAGTTAAATTAATAAACCATTCGTTTTTTTGTGATGATGAGC 1140
                                                                                                                                                                                                                                                                                                                  1141 AGAACACTAAAAGCCCATTGTTTATGGGTAAGGTTGTCAACCCCAACTCAGAAGATGTCC 1199
                                                                                                   840
                                                                                                                                                                     960
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                                                                                                                                                                                 AAGGTAAGCTTCAACATTTAGAGAATGAGTTGACTCATGACATTATTACTAAATTTTTAG
                                                                                         AGAACGAGGATCGTCGTAGCGCTTCTCGTGCACCTGCCAAAGTTAAGTATCACCGGTACTT
                                                                                                                                                AGAACGAGGATCGTCGTAGCGCTTCTCTGCACCTGCCAAAGTTAAGTATCACCGGTACTT
                                                                                                                                                                      ACGACITAAAAICIGITITIAGGCCAGITIAGGIAITACCAAAGITITITITITAACGGIGCCG
                                  /standard_name= "Ribosome binding 9.1751
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          product= "rTAP1 fusion protein"
2..1193
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       /*tag= e
/note= "TIMP-1 coding region"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                DNA sequence encoding rTAP1 fusion protein.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               note= "AAT coding region"
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'note= "linking codon"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                ABK88026 standard; DNA; 1756 BP.
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Synthetic.
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This invention relates to a novel fusion protein comprising a first protease inhibitor comprising an alphal-antitrypsin or its functionally active portion and a second protease inhibitor or its functionally active protein. The fusion proteins of the invention may act as an inhibitor of protease activity. The fusion protein of the invention is useful for inhibiting protease activity associated with a disorder such as emphysema, asthma, chronic obstructive pulmonary disease, cystic fibrosis, othis madia, officing from or a trisk for a disease or for treating an individual suffering from or a trisk for a disease or disorder involving unwanted protease activity. The proteins are useful for treating dermatological diseases such as atopic dermatitis, eczema and psoriasis, in inflammatory responses to viral infection, and for treating herpes infection, corneal or epidermal ulceration, chronic con-healing wounds, sepsis, rhemmatoid arthritis, periodontal disease, thousand and tumour metastasis and tumour angiogenesis, quartic ulceration,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          osteoporosis, Paget's disease, glomerulonephritis, scleroderma, malaria,
bacterial infection, Alzheimer's disease, hypertension and muscular
                                                                                                                                                                                                                                                                                  Novel fusion protein useful for inhibiting protease activity associated with a disorder such as emphysema, asthma, comprises a first protease inhibitor comprising alpha 1-antitrypsin and a second protease inhibitor -
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      181 TTGCCACTGCTTTCGCCATGTTGAGTTTAGGTACTAAAGCCGATACCCATGACGAGATTT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              DB 24; Length 1756;
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Score 1197.4;
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                rTAP1 fusion protein of the invention.
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    96.66;
                                                                              18-DEC-2001; 2001WO-US49256.
                                                                                                                18-DEC-2000; 2000US-256699P.
20-NOV-2001; 2001US-331966P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Best Local Similarity 99.9
Matches 1198; Conservative
                                                                                                                                                                    (ARRI-) ARRIVA PHARM INC.
                                                                                                                                                                                                       Gibson HL,
                                                                                                                                                                                                                                       2002-500631/53.
                                                                                                                                                                                                                                                         P-PSDB; AAU99889
            WO200250287-A2.
                                                27-JUN-2002
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malaria; emphysema; asthma; chronic obstructive pulmonary disease; cystic fibrosis; otitis media; otitis external; HIV; psoriasis; eczema; human immunodeficiency virus; atopic dermatitis; muscular dystrophy; herpes; ulceration; sepsis; rheumatoid arthritis; periodontal disease;
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                                                                                                                                                                                                                                                                                                                                                                                                                                                               TCTTAACTATTGATGAAAAGGGTACCGAGGCCGCGGGGGCTATGTTCCTGGAAGCTATTC 1080
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                                                                               GITTATITITGICIGAAGGITTAAAATIGGITGACAAATICCIAGAAGACGICAAGAAAC 420
            AAGTICCAAIGAAAAAGACIGGGIAIGIICAAIAIICAACAIIGCAAAAAIIAAGII 720
                                                                                                                                                                                                                      840
                                                                                                                                                                                                                                                                                                         AGAACGAGGATCGTCGTAGCGCTTCTCTGCACCTGCCAAAGTTAAGTATCACCGGTACTT 900
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                                                                                                                                                                           AAGGTAAGCTTCAACATTTAGAGAATGAGTTGACTCATGACATTATTACTAAATTTTTAG
                                         TATATCATAGTGAGGCTTTTACCGTTAATTTTGGTGATACTGAGGAAGCTAAAAGCAAA
                                                                                                                         ATCGTGATACCGTCTTCGCACTAGTTAACTATTTTTTTCAAGGGTAAGTGGGAACGTC
                                                                                                                                                                 CTTTCGAGGTTAAAGATACTGAAGAAGAAGATTTTCATGTTGATCAAGTTACTACTGTCA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Yeast; alpha factor; gene; ds; Alzheimer's disease; SLAP1;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       DNA sequence encoding SLAP1 fusion protein.
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This invention relates to a novel fusion protein comprising a first protease inhibitor comprising an alphal-antitrypsin or its functionally active portion and a second protease inhibitor or its functionally active protein. The fusion proteins of the invention may act as an antibitor of protease activity. The fusion protein of the invention is useful for inhibiting protease activity associated with a disorder such as emphysema, chronic obstructive pulmonary disease.

Cystic fibrosis, otilis media, otitis external or HIV infection, or for treating an individual suffering from or at risk for a disease or disorder involving unwanted protease activity. The proteins are useful for treating dermatological diseases such as atopic dermatitis, eczema and psociasis, in inflammatory responses to viral infection, and for treating wounds, sepsis, rheumatoid arthitis, periodomical disease, nor healing wounds, sepsis, rheumatoid arthitis, periodomical disease,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Novel fusion protein useful for inhibiting protease activity associated with a disorder such as emphysema, asthma, comprises a first protease inhibitor comprising alpha 1-antitrypsin and a second protease
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             tumour metastasis and tumour anglogenesis, gastric ulceration, osteoporosis, Paget's disease, glomerulonephritis, scleroderma, malaria, bacterial infection, Alzheimer's disease, hypertension and muscular
tumour metastasis; tumour angiogenesis; osteoporosis; Paget's disease; glomerulonephritis; scleroderma; hypertension.
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                dystrophy. The present sequence represents the DNA encoding the SLAPI fusion protein of the invention.
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99.7%; Pred. No. 1.6e-289;
11ve 0; Mismatches 4;
                                                                                                                                                                                                           'product- "SLAP1 fusion protein"
                                                                                                                                                                                                                                             c
"SLPI coding region"
                                                                                                                                                                                                                                                                                                                                                    /*tag= e
/note= "AAT coding region"
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/note= "linking codon"
336..1517
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                                                                                                      Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Pemberton P;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                         18-DEC-2001; 2001WO-US49256
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              .8-DEC-2000; 2000US-256699P
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   (ARRI-) ARRIVA PHARM INC.
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                                                   Homo sapiens.
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1 TCTAGACCATGGAAGACCCTCAAGGCGACGCCGCTCAAAAACCGACACCAGTCATCACG 60

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1405 CAATGAGCATTCCACCAGAAGTTAAATTTAATAAACCATTCGTTTTTCTGATGATCGAGC 1464

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AAGGCCATG
CCAAGACCATCCGACTTTT.
GTATAGACAATTAGCTCATC
TGCCACTGCTTTCGCCATGTT
agaaggtttaaactttaatt
AAGAGTTGTTGAGAACTTTGAA'
GTTATTTTGTCTGAAGGTTTAAAA:
TATATCATAGTGAGGCTTTTACC
TTAATGATTATGTTGAGAAAGGCA(
ATCGTGATACCGTCTTCGCACT/
CTTTCGAGGTTAAAGATACTGAAGAGGAAGATTTCAT(
aagttccaatgatgaaaagactggg
CTTGGGTCTTATTAATGAAGTV
AAGGTAAGCTTCAACATTTAGAGAATG
agaacgaggatcgtcgtagcg
ACGACTTAAAATCTGTTTTAG
ATTTGAGTGGTGTTACTGAAG!
TCTTAACTATTGATGAAAGG
CAATGAGCATTCCACCAGAAGTTAAATTTAATAAC

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This invention relates to a novel fusion protein comprising a first protease inhibitor comprising an alphal-antitrypsin or its functionally active portion and a second protease inhibitor or its functionally active protein. The fusion proteins of the invention may act as an inhibitor of protease activity. The fusion protein of the invention
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                                                                                                                                                                                                 TAP1; gene; ds; Alzheimer's disease; tumour angiogenesis; malaria; emphysema; asthma; chronic obstructive pulmonary disease; cystic fibrosis; otitis media; otitis external; HIV; psoriasis; eczema; human immunodeficiency virus; atopic dermatitis; muscular dystrophy; herpes; ulceration; sepsis; rheumatoid arthritis; periodontal disease; tumour metastasis; osteoporosis; Paget's disease; scleroderma; glomerulonephritis; hypertension.
/standard_name= "Ribosome binding site"
9..1751
                                                                                                                                                                                                                                                                                                                                                                                                                             'product= "TAP1 fusion protein"
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/note= "TIMP-1 coding region"
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/note= "AAT coding region"
                                                                                                                                                                           DNA sequence encoding TAP1 fusion protein.
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/note= "linking codon"
567..1748
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                                                                                              ABK88023 standard; DNA; 1756 BP.
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20-NOV-2001; 2001US-331966P.
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ts useful for inhibiting protease activity associated with a disorder such as emphysema, asthma, chronic obstructive pulmonary disease, creating as emphysema, asthma, chronic obstructive pulmonary disease, cystic fibrosis, citis media, citis external or HIV infection, or for treating an individual suffering from or at risk for a disease or disorder involving unwanted protease activity. The proteins are useful for treating dermatological diseases such as atopic dermatitis, eccemna and psoriasis, in inflammatory responses to viral infection, chronic non-healing wounds, sepsis, rheumatorid arthritis, periodontal disease, tumour metastasis and tumour anglogenesis, gastric ulceration, costeoporosis, Paget's disease, glomerulonephritis, scleroderma, malaria, bacterial infection, Alzhelmer's disease, hypertension and muscular dystrophy. The present sequence represents the DNA encoding the content of the invention.
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                                                                          ACCATCCGACTTTAATAAAATTACTCCAAATTTAGCCGAATTTGCTTTTTCTTTGTATA 126
                                                                                                                   CTGCTTTCGCCATGTTGAGTTTAGGTACTAAAGCCGATACCCATGACGAGATTTTAGAAG 246
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                                                              7 CCATGGAAGACCCTCAAGGCGACGCCGCTCAAAAAACCGACACCAGTCATCACGACCAAG 66
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                    78.0%; Score 1188.8; DB 24; Length 1756; 99.8%; Pred, No. 8.3e-289;
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                                          2; Indels
Sequence 1756 BP; 493 A; 395 C; 373 G; 495 T; 0 other;
                                         0; Mismatches
                             Best Local Similarity 99.8 Matches 1190, Conservative
                      Query Match
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NTAP1; gene; ds; Alzheimer's disease; tumour angiogenesis; malaria; emphysema; asthma; chronic obstructive pulmonary disease; cystic fibrosis; ottitis media; ottitis external; HIV; psoriasis; eczema; human immunodeficiency virus; atopic dermatitis; muscular dystrophy; herpes; ulceration; sepsis; rheumatoid arthritis; periodontal disease; tumour metastasis; osteoporosis; Paget's disease; scleroderma;
                                                                                                                                                                                         GTGGTGTTACTGAAGAAGCTCCATTAAAATTGAGTAAAGCTGTTCACAAAGCCGTCTTAA 1026
                                                                                                                                             CTATTGATGAAAAGGGTACCGAGGCCGCCGCGCTATGTTCCTGGAAGCTATTCCAATGA 1086
                                                                                                                                                                                                                        GCATTCCACCAGAAGTTAAATTTAATAAACCATTCGTTTTTTCTGATGATCGAGCAGAACA 1146
                                                                                                                                                                                                                                   1642 GCATTCCACCAGAAGTTAAAATTTAAAACCATTCGTTTTTTCTGATGATGGAGGAGAAGA 1701
                                                                                      AGCTTCAACATTTAGAGAATGAGTTGACTCATGACATTATTACTAAATTTTTAGAGAACG
                                            AGGATCGTCGTAGCGCTTCTCTGTACCTGCCAAAGTTAAGTATCACCGGTACTTACGACT
                                                                                                                                                                                                                                                                 site"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           /standard_name= "Ribosome binding
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           "NTAP1 fusion protein
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/note= "TIMP-1 coding region"
                                                                                                                                                                                                                                                                                                                                                                                                               DNA sequence encoding N-TAP1 fusion protein.
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/note= "linking codon"
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                                                                                                                                      δ
                                                                                                                                                                                                                                                                                                                                                                                               This invention relates to a novel fusion protein comprising a first protease inhibitor comprising an alphal-antitrypain or its functionally active portion and a second protease inhibitor or its functionally active protein. The fusion protease activity of the invention may act as an inhibitor of protease activity. The fusion protein of the invention is useful for inhibiting protease activity associated with a disorder such as emphysema, asthma, chronic obstructive pulmonary disease, cystic fibrosis, otitis media otitis external or HIV infection, or for treating an individual suffering from or at risk for a disease or city treating dermatological diseases such as atopic dermatitis, eczema and psoriasis, in inflammatory responses to viral infection, and for treating herpes infection, corneal or epidermal ulceration, chronic treating wounds, sepsis, rheumatoid arthritis, periodontal disease, tumour metastasis and tumour angiogenesis, gastric ulceration, malaxia
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               ó
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    osteoporosis, Paget's disease, glomerulonephritis, scleroderma, malaria, bacterial infection, Alzheimer's disease, hypertension and muscular dystrophy. The present sequence represents the DNA encoding the NTAPI fusion protein of the invention.
                                                                                                                                                                                                                               Novel fusion protein useful for inhibiting protease activity associated with a disorder such as emphysema, asthma, comprises a first protease inhibitor comprising alpha 1-antitrypsin and a second protease
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ATAGACAATTAGCICAICAAAGTAATICTACTAACATTTTTTTTTAGTCCTGTTTCTATTG 183
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  243
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          AGTIGITGAGAACITIGAATCAACCIGATICICAATIGCAATIAACIACIGGIAACGGIT 363
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           63
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   CCACTGCTTTCGCCATGTTGAGTTTAGGTACTAAAGCCGATACCCATGACGAGATTTTAG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AAGGTTTAAACTTTAATTTGACCGAAATCCCAGAAGCCCCAAATTCACGAGGGTTTTCAAG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            685 AGTIGITGAGAACTITGAATCAACCIGATTCTCAATTGCAATTAACTACTGGTAACGGTT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         TATITITGTCTGAAGGTTTAAAATTGGTTGACAAATTCCTAGAAGACGTCAAGAAACTAT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           4 AGACCATGGAAGACCCTCAAGGCGACGCCGCTCAAAAAACCGACACCAGTCATCACGACC
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Sequence 1582 BP; 464 A; 333 C; 329 G; 456 T; 0 other;
                                                                                                                                                                                                                                                                                                                                                         Example 2; Page 85-86; 134pp; English.
                                                                                                                  Pemberton P;
2000US-256699P.
2001US-331966P.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Matches 1190; Conservative
                                                                   (ARRI-) ARRIVA PHARM INC
                                                                                                                    Gibson HL,
                                                                                                                                                                    2002-500631/53
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                                                                                                                                                                                             P-PSDB; AAU99883
18-DEC-2000;
20-NOV-2001;
                                                                                                                                                                                                                                                                                                               inhibitor
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                                                                                                                       Barr PJ,
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Alpha-1-antitrypsin; AAT; human; gene; ds; protease inhibitor; malaria; emphysema; asthma; chronic obstructive pulmonary disease; eczema; cystic fibrosis; otitis media; otitis external; HIV; psoriaais; human immunodeficiency virus; atopic dermatitis; muscular dystrophy; herpes; ulceration; sepsis; rheumatoid arthritis; periodontal disease; tumour metastasis; tumour angiogenesis; osteoporosis; Paget's disease; glomerulonephritis; scleroderma; Alzheimer's disease; hypertension.
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        1084 TGAGCATTCCACCAGAAGTTAAATTTAATAAACCATTCGTTTTTCTGATGATCGAGCAGA 1143
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  TGAGTGGTGTTACTGAAGAAGCTCCATTAAAATTGAGTAAAAGCTGTTCACAAAGCCGTCT 1023
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              TAACTATTGATGAAAAGGGTACCGAGGCCGCCGGCGCTATGTTCCTGGAAGCTATTCCAA 1083
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          1165 GTAAGCTTCAACATTTAGAGAATGAGTTGACTCATGACATTATTACTAAATTTTTAGAGA 1224
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ACGAGGATCGTCGTAGCGCTTCTCTGCACCTGCCAAAGTTAAGTATCACCGGTACTTACG 1284
                                                                                                                                                                                                              GTAAGCTICAACAITIAGAGAATGAGTIGACTCATGACATTATTACTAAATTITTAGAGA 843
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                844 ACGAGGAICGICGIAGCGCTICICGCACCIGCCAAAGITAAGIATCACCGGIACITACG 903
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    1405 TAACTATTGATGAAAAGGTACCGAGCCGCCGCGCGCTATGTTCCTGGAAGCTATTCCAA
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                                                        GTGATACCGTCTTCGCACTAGTTAACTATATTTTTTCAAGGGTAAGTGGGAACGTCCTT
                                                                                                                                    TCGAGGTTAAAAGATACTGAAGAGGAAGATTTTCATGTTGATCAAGTTACTACTGTCAAAG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ACTITAAAATCTGTTTTAGGCCAGTTAGGTATTACCAAAGTTTTTTCTAACGGTGCCGATT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  1144 ACACTAAAAGCCCATTGTTTATGGGTAAGGTTGTCAACCCAACTCAGAAGATGTC 1198
805 ATCATAGTGAGGCTTTTACCGTTAATTTTGGTGATACTGAGGAAGCTAAAAAGCAAATTA
                                      ATGATTATGTTGAGAAAGGCACCCAGGGTAAGATCGTTGACCTAGTTAAAGAATTAGATC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          1525 ACACTAAAAGCCCATTGTTATGGGTAAGGTTGTCAACCCAACTCAGAAGTAGTC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          DNA encoding human alpha-1-antitrypsin (AAT) protein.
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/note= "No start or stop codon shown"
/product= "Alpha-1-antitrypsin"
                           /partial
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WO200250287-A2

27-JUN-2002

18-DEC-2001; 2001WO-US49256

18-DEC-2000; 2000US-256699P. 20-NOV-2001; 2001US-331966P.

(ARRI-) ARRIVA PHARM INC

Pemberton P; Glbson HL, Barr PJ,

WPI; 2002-500631/53 P-PSDB; AAU99873 Novel fusion protein useful for inhibiting protease activity associated with a disorder such as emphysema, asthma, comprises a first protease inhibitor comprising alpha 1-antitrypsin and a second protease

Disclosure; Page 24-25; 134pp; English.

This invention relates to a novel fusion protein comprising a first
protease inhibitor comprising an alphal-antitrypsin or its functionally
active portion and a second protease inhibitor or its functionally
active protein. The fusion proteins of the invention may act as an
cutive protein. The fusion proteins of the invention any act as an
cutive protein. The fusion proteins of the invention
is useful for inhibiting protease activity associated with a disorder
cystic fibrosis, othis media, oritis serverial or HIV infection, or
for treating an individual suffering from or at risk for a disease or
disorder involving unwanted protease activity. The proteins are useful
for treating dermachlogical diseases and a stopic dermatitis, eczema
and sporiasis, in inflammatory responses to viral infection, and for
treating herpes infection, corneal or epidermal ulceration, chronic
non-healing wounds, sepsis, rheumatoid arthritis, periodontal disease,
tumour metastasis and tumour anglogenesis, gastric ulceration,
bacterial infection, Alzheimer's disease, hypertension and muscular
dystrophy. The present sequence represents the DNA encoding the human
a for the invention protein used to create the fusion protein of the invention.

Sequence 1182 BP; 369 A; 214 C; 229 G; 370 T; 0 other;

ö 180 TTCGCCATGTTGAGTTTAGGTACTAAAGCCGATACCCATGACGAGATTTTAGAAGGTTTA 251 AACTITAATTTGACCGAAATCCCAGAAGCCCAAATTCACGAGGGTTTTCAAGAGTTGTTG 311 131 132 TTAGCTCATCAAAGTAATTCTACTAACATTTTTTTAGTCCTGTTTCTATTGCCACTGCT 191 Gaps 121 TTAGCTCATCAAAGTAATTCTACTAACATTTTTTTTAGTCCTGTTTCTATTGCCACTGCT 12 GAAGACCCTCAAGGCGACGCCGCTCAAAAACCGACACCAGTCATCACGACCAAGACCAT 72 CCGACTTTTAATAAAATTACTCCAAATTTAGCCGAATTTGCTTTTTCTTTGTATAGACAA ö Length 1182; Indels 77.5%; Score 1182; DB 24; 100.0%; Pred. No. 3.7e-287; tive 0; Mismatches 0; Matches 1182; Conservative Best Local Similarity Query Match 192 252 유 ò 8 음 ð ò ò g

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CCACCAGAAGTTAAATTTAATAAACCATTCGTTTTTCTGATGATCGAGCAGAACACTAAA 1151
                                241 AACTTTAATTTGACCGAAATCCCAGAAGCCCAAATTCACGAGGGTTTTCAAGAGTTGTTG 300
                                                                        GICTICGCACTAGITAACTATATITITCAAGGGTAAGIGGGAACGICCTITCGAGGIT 611
                                                                                                                                                                                                                                                                         ATGAAAAGACTGGGTATGTTCAATATTCAACATTGCAAAAATTAAGTTCTTGGGTCTTA 731
                                                                                                                                                                                                                                                                                             661 ATGAAAAGACTGGGTATGTTCAATATTCAACATTGCAAAAAATTAAGTTCTTGGGTCTTA 720
                                                                                                                                                                                                                                                                                                                            971
                    AGAACTTTGAATCAACCTGATTCTCAATTGCAATTAACTACTGGTAACGGTTTATTTTG 371
                                                              TCTGAAGGTTTAAAATTGGTTGACAAATTCCTAGAAGACGTCAAGAAACTATATCATAGT 431
                                                                                                      GAGGCTTTTACCGTTAATTTTGGTGATACTGAGGAAGCTAAAAAGCAAATTAATGATTAT 491
                                                                                                                                                                                                                                                                                                                 791
                                                                                                                 1032 GATGAAAAGGGTACCGAGGCCGCCGCGCGTATGTTCCTGGAAGCTATTCCAATGAGCATT
                                                                                                                                               GTTGAGAAAGGCACCCAGGGTAAGATCGTTGACCTAGTTAAAGAATTAGATCGTGATACC
                                                                                                                                                                                                            541 GTCTTCGCACTAGTTAACTATTTTTTTCAAGGGTAAGTGGGAACGTCCTTTCGAGGTT
                                                                                                                                                                                                                                                                                                                 CAACATTTAGAGAATGAGTTGACTCATGACATTATTACTAAATTTTTAGAGAACGAGGAT
                                                                                                                                                                                                                                                                                                                                                                                                                                           972 GTIACTGAAGAAGCTCCATTAAAATTGAGTAAAGCTGTTCACAAAGCCGTCTTAACTATT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Codon-optimised RAmy3D signal fused to DNA encoding mature AAT.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       1152 AGCCCATTGTTTATGGGTAAGGTTGTCAACCCAACTCAGAAG 1193
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to exemplify the invention. The specification describes a method for producing mature alphal-antitypsin (AAT). The protein is used to producing mature heterologous protein in monocotyledonous plant cells. The method comprises transforming the cells with a chimeric gene comprising a monocotyledon transcription regulator, inducible either during seed maturation or by adding/removing a small molecule, DNA encoding the heterologous protein, and DNA encoding a signal peptide, with the signal peptide causing secretion of the protein from the cell. The proteins expressed in this manner include mature glycosylated alpha proteins expressed in this manner include mature glycosylated alpha increases its serum half-life, mature glycosylated antithrombin III (ATIII), mature human serum albumin (HSA) having the native folding pattern as shown by bilirubin-binding characteristics, or mature active cubtilisin BPN'. These proteins are useful therapeutically (e.g. AAT for treating emphysema, ATIII as antithrombotic and HSA as blood replacement) or as industrial enzymes (BPN' is used in detergents).
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               present sequence encodes a fusion protein of codon-optimised RAmy3D
Protein expression; monocotyledon plant cell; glycosylated alpha 1-antitrypsin; AAT; glycosylated alpha 1-antitrypsin; AAT; glycosylated artithrombin III; ATIII; human serum albumin; HSA; subtilisin BBN'; treatment; emphysema; antithrombotic; blood replacement; ss.
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        132 TTAGCTCATCAAAGTAATTCTACTAACATTTTTTTTAGTCCTGTTTCTATTGCCACTGCT 191
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Gaps
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Expressing mature, glycosylated proteins in monocotyledonous plant cells - from chimeric gene including signal peptide sequence, specifically therapeutic agents and industrial enzymes
                                                                                                                                                                                                                                                                                              l..75
/*tag= a
/note= "codon-optimised RAmy3D signal sequence"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Score 628.4; DB 19; Length 1260; Pred. No. 5.4e-148;
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/note= "encodes mature AAT"
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                                                                                                                                                                                                                                                                Location/Qualifiers
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97US-0038168.
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Best Local Similarity
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976 AGCGTGCTGGGCCAGCTGGGCATCACGAAGGTCTTCAGCAACGGCGGGGGGGCTCTCCGGC 1035 972 GTTACTGAAGAAGCTCCATTAAAATTGAGTAAAGCTGTTCACAAAGCCGTCTTAACTATT 1031 1032 GATGAAAAGGGTACCGAGGCCGCCGCCGTATGTTCCTGGAAGCTATTCCAATGAGCATT 1091 1092 CCACCAGAAGTTAAATTAATAAACCATTCGTTTTTCTGATGATCGAGCAGAACACTAAA 1151 611 675 671 735 791 672 ATGAAAAGACTGGGTATGTTCAATATTCAACATTGCAAAAAATTAAGTTCTTGGGTCTTA 731 491 551 196 CTCGCGCCACCAGTCCAACTCCAACATCTTCTTCAGCCCGGTGAGCATCGCCACCGCC 255 AACTITAATTTGACCGAAATCCCCAGAAGCCCCAAATTCACGAGGGTTTTCAAGAGTTGTTG 311 372 TCTGAAGGTTTAAAATTGGTTGACAAATTCCTAGAAGACGTCAAGAAACTATATCATAGT 431 GICTICGCACIAGITAACIATITITITICAAGGGTAAGIGGGAACGICCITICGAGGIT 1096 GACGAGAAGGGACGGAAGCTGCCGGGGCCATGTTCCTGGAGGCCATCCCCATGTCCATC TICGCCATGITGAGITITAGGIACIAAAGCCGAIACCCAIGACGAGAITITAGAAGGITIA AGAACTTTGAATCAACCTGATTCTCAATTGCAATTAACTACTGGTAACGGTTTATTTTG GAGGCTTTTACCGTTAATTTTGGTGATACTGAGGAAGCTAAAAAAGCAAATTAATGATTAT GTTGAGAAAGGCACCCAGGGTAAGATCGTTGACCTAGTTAAAGAATTAGATCGTGATACC 556 GTCGAGAAGGGGACCCAGGGCAAGATCGTGGACCTGGTCAAGGAATTGGACAGGGACACC 612 AAAGATACTGAAGAGGAAGATTTTCATGTTGATCAAGTTACTACTGTCAAAGTTCCAATG 792 CAACATTTAGAGAATGAGTTGACTCATGACATTATTACTAAATTTTTAGAGAACGAGGAT 852 CGTCGTAGCGCTTCTCTGCACCTGCCAAAGTTAAGTATCACCGGTACTTACGACTTAAAA 1036 GTGACGGAGGAGGCCCCCTGAAGCTCTCCAAGGCCGTGCACAAGGCGGTGCTCACGATC 1216 AGCCCCTCTTCATGGGGAAGGTCGTCAACCCCACGCAGAAG 1257 1152 AGCCCATTGTTTATGGGTAAGGTTGTCAACCCAACTCAGAAG 1193

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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             192 TTCGCCATGTTGAGTTTAGGTACTAAAGCCGATACCCATGACGAGATTTTAGAAGGTTTA 251
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        0; Gaps
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        12 GAAGACCCTCAAGGCGACGCCGCTCAAAAACCGACACCAGTCATCACGACCAAGACCAT 71
                                                                                                                                                                                                                                                                                                                                                                                                                 The sequence of a human alpha-1-antitrypsin cDNA clone is given in AAQ89254. Expression of the cDNA in host cell transformants has allowed production of recombinant alpha-1-antitrypsin.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             220 CTGGCACACCAGTCCAACAGCACCAATATCTTCTTCTCCCCAGTGAGCATCGCTACAGCC
                                                                                                                                                                                                                                                                                                                                                                         Human alphal-antitrypsin (al-AT) cDNA sequence - can be used for
                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Query Match 28.3%; Score 432.2; DB 16; Length 1312; Best Local Similarity 60.4%; Pred. No. 1.1e-98; Matches 713; Conservative 0; Mismatches 468; Indels 0;
                                                                                                                                                                                                                                                                                                                                                                                                                                                   Sequence 1312 BP; 339 A; 368 C; 324 G; 281 T; 0 other;
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                                                                  Alpha-1-trypsin; protease-inhibitor; ss
                                                                                                    Cocation/Qualifiers
28..1258
AAQ89254 standard; cDNA; 1312 BP
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8705-0133190

8705-0133190

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9205-099556
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                                                  Human alpha-1-trypsin cDNA
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28..99
/*tag= b
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/*tag= c
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                                                                                                                                                                                                                                                                                                                                                                                  the expression of al-AT
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                                                                                   Homo sapiens
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18-NOV-1992;
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03-MAR-1987
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22-AUG-1989
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1000 AGCGTCCTAGGTCAACTGGGCATCACTAAGGTCTTCAGCAATGGGGCTGACCTCTCCGGG 1059
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        1092 CCACCAGAAGTTAAATAAACCATTCGTTTTTCTGATGATCGAGCAGAACACTAAA 1151
                           252 AACTITAATITGACCGAAATCCCAGAAGCCCAAATTCACGAGGGTTTTCAAGAGTTGTTG 311
                                                                                                    312 AGAACTTTGAATCAACCTGATTCTCAATTGCAATTAACTACTGGTAACGGTTTATTTTTG 371
                                                                                                                                                      400 CGTACCCTCAACCAGCCAGACAGCCAGCTCCAGCTGACCACCGGCAATGGCCTGTTCCTC 459
                                                                                                                                                                                                            372 TCTGAAGGTTTAAAATTGGTTGACAAATTCCTAGAAGACGTCAAGAACTATATCATAGT 431
                                                                                                                                                                                                                                                              460 AGCGAGGCCTGAAGCTAGTGGATAAGTTTTTGGAGGATGTTAAAAAGTTGTACCACTCA 519
                                                                                                                                                                                                                                                                                                                                                                                                                        GTTGAGAAAGGCACCCAGGGTAAGATCGTTGACCTAGTTAAAGAATTAGATCGTGATACC 551
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                792 CAACATTTAGAGAATGAGTTGACTCATGACATTATTACTAAATTTTTAGAGAACGAGGAT 851
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       852 CGTCGTAGCGCTTCTCTGCACCTGCCAAAGTTAAGTATCACCGGTACTTACGACTTAAAA 911
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               912 TCTGTTTTAGGCCAGTTAGGTATTACCAAAGTTTTTTCTAACGGTGCCGATTTGAGTGGT 971
                                                                                                                                                                                                                                                                                                                432 GAGGCTTTTACCGTTAATTTTGGTGATACTGAGGAAGCTAAAAAGCAAATTAATGATTAT 491
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   CCGACTTTTAATAAAATTACTCCAAATTTAGCCGAATTTGCTTTTTCTTTGTATAGACAA 131
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              160 CCAACCTTCAACAAGATCACCCCCAACTTGGCTGAGTTCGCCTTCAGCCTATACCGCCAG 219
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                                Human alpha-1-antitrypsin; ATR-1; antibody; ATR-1 deficiency; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  28.3%; Score 432.2; DB 19; Length 1312; ilarity 60.4%; Pred. No. 1.1e-98; Conservative 0; Mismatches 468; Indels 0;
                                                                                                                                                                                                                                                                                                                                                                                             encoding alpha-1 anti-trypsin - useful for, e.g. producing
                                                                                                                                                                                                                                                                                                                                                                                                                                                 This is the nucleotide sequence encoding the novel human
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sequence 1312 BP; 339 A; 368 C; 324 G; 281 T; 0 other;
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                                                                                                  /*tag= a
/product= "alpha-1-antitrypsin"
            Nucleotide sequence of the alpha-1-antitrypsin.
                                                                                                                                                                                                                                                                                                                                         Thirumalachary C,
                                                                           Location/Qualifiers
                                                                                                                                                                                                                                                                                                                                                                                                        recombinant alpha-1 anti-trypsin
                                                                                                                                                                                                                                                                                                                                                                                                                              Claim 1; Fig 1; 15pp; English.
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8405-0638980.
8705-0022543.
8705-0133190.
8805-0246912.
8905-0398288.
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93US-0086442.
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P-PSDB; AAW56709.
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15-DEC-1987;
16-SEP-1988;
                                                        Homo sapiens
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02-JUL-1993;
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1180 CGCCCCGAGGTCCAAGTTCAACAACCCTTTGTCTTCTTAATGATTGAACAAAATACCAAG 1239
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        1032 GATGAAAAGGGTACCGAGGCCGCCGCGCGTATGTTCCTGGAAGCTATTCCAATGAGCATT 1091
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          1092 CCACCAGAAGTTAAATTTAATAAACCATTCGTTTTTCTGATGATCGAGCAGAACACTAAA 1151
                                                                                         971
640 GIIIIIIGCICIGGIGAAIIACAICIICITIAAAGGCAAAIGGGAGAGACCCIIIGAAGIC 699
                                                                                                                                                                                                                                                                                                                                                                                                                                                                   671
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            940 AGAAGGICIGCCAGCITACATITACCCAAACIGICCATTACIGGAACCTAIGAICIGAAG 999
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               851
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  672 AIGAAAAGACIGGGIAIGIICAAIAIICAACAIIGCAAAAAAIIAAGIICIIGGGICIIA 731
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Alphal-antitrypsin; neutrophil elastase inhibitor; human; ss; chronic obstructive pulmonary emphysema; infantile liver cirrhosis.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       580 GTGGAGAAGGGTACTCAAGGGAAAATTGTGGATTTGGTCAAGGAGCTTGACAGAGACACA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          372 ICTGAAGGTTTAAAATTGGTTGACAAATTCCTAGAAGACGTCAAGAAACTATATCATAGT
                                                                                                                                                                                                                                                                     GITGAGAAAGGCACCCAGGGTAAGATCGTTGACCTAGTTAAAGAATTAGATCGTGATACC
                                                                                                                                                                                                                                                                                                                                                                    552 GTCTTCGCACTAGTTAACTATATTTTTTCAAGGGTAAGTGGGAACGTCCTTTCGAGGTT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                   612 AAAGATACTGAAGAGGAAGATTTTCATGTTGATCAAGTTACTACTGTCAAAGTTCCAATG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  792 CAACATTTAGAGAATGAGTTGACTCATGACATTATTACTAAATTTTTAGAGAACGAGGAT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 852 CGTCGTAGCGCTTCTCTGCACCTGCCAAAGTTAAGTATCACCGGTACTTACGACTTAAAA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                912 TCTGTTTTAGGCCAGTTAGGTATTACCAAAGTTTTTTCTAACGGTGCCGATTTGAGTGGT
                                                                                                                                                                    GAGGCTTTTTACCGTTAATTTTGGTGATACTGAGGAAGCTAAAAAGCAAATTAATGATTAT
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                                                                                                                                                                                                                                                                                                                                                                                                                        This sequence represents the human alphal-antitrypsin nucleotide sequence. Alphal-antitrypsin is an important protease inhibitor, the major function of which is to inhibit neutrophil elastess. Low levels of alphal-antitrypsin in the blood are associated with chronic obstructive pulmonary emphysema and infantile liver cirrhosis. A vector comprising a mammalian alphal-antitrypsin DNA sequence that hybridises to human alphal-antitrypsin can be introduced into a host cell in a method for the production of alphal-antitrypsin.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  CCGACTITIAATAAAATTACTCCAAATTTAGCCGAATTTGCTTTTTCTTTGTATAGACAA 131
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     TTTGCAATGCTCTCCCTGGGGACCAAGGCTGACACTCACGATGAATCCTGGAGGCCTG 339
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                CCAACCTTCAACAAGATCACCCCCAACTTGGCTGAGTTCGCCTTCAGCCTATACCGCCAG 219
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             CTGGCACACCAGTCCAACAGCACCAATATCTTCTTCTCCCCAGTGAGCATCGCTACAGCC 279
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      TTCGCCATGTTGAGTTTAGGTACTAAAGCCGATACCCATGACGAGATTTTAGAAGGTTTA 251
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       AACTTTAATTTGACCGAAATCCCAGAAGCCCAAATTCACGAGGGTTTTCAAGAGTTGTTG 311
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AGAACTITGAATCAACCTGATTCTCAATTGCAATTAACTACTGGTAACGGTTTATTTTG 371
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       400 CGTACCCTCAACCAGCCAGACAGCTCCAGCTGACCACCGGCAATGGCCTGTTCCTC 459
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                GAAGACCCTCAAGGCGACGCCGCTCAAAAACCGGACACCAGTCATCACGACCAAGACCAT 71
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    132 TTAGCTCATCAAAGTAATTCTACTAACATTTTTTTTAGTCCTGTTTCTATTGCCACTGCT
                                                                                                                                                                                                                                                                                                                                    Preparing alphal-antitrypsin for inhibiting neutrophil elastase
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   DB 21; Length 1312;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ö
                                                                                                                                                                                                                                                                                                                                                involves transfecting host cell with vector comprising alphal-antitrypsin DNA sequence that hybridizes to human alphal-antitrypsin CDNA, or its complement
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           468; Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Sequence 1312 BP; 339 A; 368 C; 324 G; 281 T; 0 other;
                                                                                                                                                                                                                                                                          Davie
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Score 432.2; DB 2
Pred. No. 1.1e-98;
0; Mismatches 468
                                                                                                                                                                                                                                                                        Woo SLC, Thirumalachary C, Kurachi K,
                                                                                                                                                                                                                                                                                                                                                                                                   Claim 1; Fig 1; 16pp; English.
                                                                                                                                              87US-0022543.
87US-0133190.
88US-0246912.
                                                                                                                                                                                 89US-0398288.
91US-0666450.
92US-0979556.
                                                                                                                                                                                                                                                (WASH-) WASHINGTON RES FOUND
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 28.3%;
                                                                                                                        82US-0380810.
84US-0638980.
                                                                                   98US-0009581
                                                                                                           95US-0479545
                                                                                                                                                                                                                       93US-0086442
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Local Similarity
                                                                                                                                                                                                                                                                                                             P-PSDB; AAY78890
           Homo sapiens
                                                                                   20-JAN-1998;
                                  US6025161-A.
                                                                                                                       20-MAY-1982;
07-FEB-1984;
                                                            15-FEB-2000
                                                                                                                                                          15-DEC-1987
16-SEP-1988
                                                                                                                                                                                   22-AUG-1989
11-MAR-1991
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1060 GTCACAGAGGAGGCACCCCTGAAGCTCTCCAAGGCCGTGCATAAGGCTGTGCTGACCATC 1119
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             GTTACTGAAGAAGCTCCATTAAAATTGAGTAAAGCTGTTCACAAAGCCGTCTTAACTATT 1031
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       GATGAAAAGGGTACCGAGGCCGCCGCGCTATGTTCCTGGAAGCTATTCCAATGAGCATT 1091
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                                                                                                             491
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                                                      460 AGCGAGGCCTGAAGCTAGTGGATAAGTTTTTGGAGGATGTTAAAAAGTTGTACCACTCA 519
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ischaemia reperfusion injury; haematopolesis; cancer; neuropathy; transgentc animal; Alzheimer's disease; Parkinson's disease; burn; amyotrophic lateral sclerosis; platelet disorder; thrombocytopenia; ulcer; osteoporosis; bone degenerative disorder; periodontal disease; gut protection; lung; liver fibrosis; immune deficiency; infection;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Human; secreted protein; arthritis; Crohn's disease; sepsis; shock;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                         852 CGTCGTAGCGCTTCTCTGCACCTGCCAAAGTTAAGTATCACCGGTACTTACGACTTAAAA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           940 AGAAGGTCTGCCAGCTTACATTTACCCAAAACTGTCCATTACTGGAACCTATGATCTGAAG
372 TCTGAAGGTTTAAAATTGGTTGACAAATTCCTAGAAGACGTCAAGAAACTATATCATAGT
                                                                                                          432 GAGGCTTTTACCGTTAATTTTGGTGATACTGAGGAAGCTAAAAAGCAAATTAATGATTAT
                                                                                                                                                                 520 GAAGCCTTCACTGTCAACTTCGGGGACACGGAAGAGGCCAAGAAACAGATCAACGATTAC
                                                                                                                                                                                                                    492 GTTGAGAAAGGCACCCAGGGTAAGATCGTTGACCTAGTTAAAGAATTAGATCGTGATACC
                                                                                                                                                                                                                                                       640 GITITIGCICIGGIGAAITACAICTICITIAAAGGCAAAIGGGAGAGACCCITIGAAGIC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   672 ATGAAAAGACTGGGTATGTTCAATATTCAACATTGCAAAAAATTAAGTTCTTGGGTCTTA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 792 CAACATTTAGAGAATGAGTTGACTCATGACATTATTACTAAATTTTTAGAGAACGAGGAT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               880 CAGCACCTGGAAAATGAACTCACCCAGGATATCATCACCAAGTTCCTGGAAAATGAAGAC
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                                                                                                                                                                                                                                                                                                                                                                                                                                         612 AAAGATACTGAAGAAGATTTTCATGTTGATCAAGTTACTACTGTCAAAGTTCCAATG
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            1152 AGCCCATTGTTTATGGGTAAGGTTGTCAACCCCAACTCAGAA 1192
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105 GAGGATCCCCAGGGAGATGCTGCCCAGAAGACAGATACATCCCCACCATGATCAGGATCAC 164

P δŏ

12 GAAGACCCTCAAGGCGACGCCGCTCAAAAACCGACACCAGTCATCACGACCAAGACCAT 71

251

285 ITTGCAATGCTCTCCCTGGGGACCAAGGCTGACACTCACGATGAAATCCTGGAGGGCCTG 344

TICGCCATGITGAGITITAGGIACIAAAGCCGATACCCAIGACGAGAITITAGAAGGITIA

192

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225 CIGGCACACCAGTCCAACAGCACCAATAICTICTICTCCCCAGTGAGCATCGCTACAGCC

132 TTAGCTCATCAAAGTAATTCTACTAACATTTTTTTTTAGTCCTGTTTCTATTGCCACTGCT

ŏ g δ 345 AATTICAACCICACGGAGATICCGGAGGCICAGATCCAIGAAGGCIICCAGGAACICCIC 404

AGAACTITGAATCAACCTGATTCTCAATTGCAATTAACTACTGGTAACGGTTTATTTTG 371

312

AACTTTAATTTGACCGAAATCCCAGAAGCCCAAATTCACGAGGGTTTTCAAGAGTTGTTG 311

464

524

TCTGAAGGTTTAAAATTGGTTGACAAATTCCTAGAAGACGTCAAGAAACTATATCATAGT 431

405 CGTACCCTCAACCAGCCAGACAGCCAGCTCCAGCTGACCACCGGCAATGGCCTGTTCCTC

GAGGCTTTTACCGTTAATTTTGGTGAT-ACTGAGGAAGCTAAAAAAGCAAATTAATGATTA 490 GAAGCCTTCACTGTCAACTTCGGGGATCACCGAAGAGGCCCAAGAAACAGATCAACGATTA 584

432 525 491

07-MAR-2000; 13-SEP-2001 Zhao QA, Tang YT,

continuous such as arthritis, neptritis, Cronn's disease, isohaemia-reperfusion injury, shock, sepsis, immune responses, and is involved in increasing haematopolesis, stem cell survival, bone growth and remodeling, (1), (11) and modulators of (11) are useful for prophylaxis or treatment of one or more cancers. (11) is also useful for creating transgenic animals useful for studying the in vivo activities of the polypeptide as well as for studying modulators of the polypeptides. (1) induces the proliferation of neural cells and regeneration of nerve and brain tissue and is useful for the treatment of central and periods the proliferation of neural cells and regeneration of nerve and brain tissue and is useful for the treatment of central and periods. In addition, (1) is involved in chemotactic or chemokinetic activity, regulation of haematopoiesis and is useful for treating myeloid or lymphoid cell disorders platelet disorders such as thrombocytopenia and for regeneration of bone, cartilage, tendon, ligament and/or nerve tissue growth, and in tissue repair, healing of burns, incisions, culcars, for treating osteoporosis, osteoarthritis, bone degeneration or egeneration and treatment of lung or liver fibrosis, reperfusion injury in various tissues, various immune deficiencies and disorders including severe combined immunodeliciency (SCID), bacterial or fungal infections, autoimmune disorders of multiple sclerosis.

Creatility, metabolism, catabolism, anabolism, storage or elimination of deterny fat, libid, protein, and an electric composition to raise an antigen in a vaccine composition to raise an antigen in a vaccine composition to raise an entigen in a vaccine composition to raise an ording accined and protein reducing effects, immunos electric produces of the invention. The invention relates to novel isolated human secreted polypeptides (I) and polynucleotides (II). (I) and (II) are useful for treating inflammatory conditions such as arthritis, nephritis, Crohn's disease, Zhou P; severe combined immunodeficiency; SCID; autoimmune disorder; allergy; multiple sclerosis; rheumatoid arthritis; diabetes mellitus; asthma; fertility; analgesic; pain; antigen; ss. Novel polypeptides and nucleic acids obtained from cDNA libraries prepared from various human tissues, for diagnosis and treatment of cancer, neurological, inflammatory, and autoimmune disorders -C, Wehrman T, Ren F, Ma Y, Zhou Zhang J, Chen R, Xue AJ, Wang J; Sequence 1367 BP; 357 A; 392 C; 323 G; 295 T; 0 other; Claim 1; SEQ ID No 133; 107pp; English. χu C, of the invention. Drmanac RT, 19-MAY-2000; 2000US-0574454. 17-JUN-2000; 2000US-0596193. 14-JUL-2000; 2000US-0616847. 19-SEP-2000; 2000US-0665363. 28.1%; Liu C, Asundi V, 2000US-0519705 05-MAR-2001; 2001WO-US04942 20-OCT-2000; 2000US-0693267 WPI; 2001-589934/66. Yang Y, (HYSE-) HYSEQ INC. P-PSDB; AAU28152. WO200166689-A2.

1091 TCCACCAGAAGTTAAATTTAATAAACCATTCGTTTTTCTGATGATCGAGCAGAACACTAA 1150 1031 TGATGAAAAGGGTACCGAGGCCGCCGGCGTATGTTCCTGGAAGCTATTCCAATGAGCAT 1090 1065 GGTCACAGAGGAGGCACCCCTGAAGCTCTCCAAGGCCGTGCATAAGGCTGTGCTGACCAT 1125 CGACGAGAAGGGGACTGAAGCTGCTGGGGGCCATGTTTTAGAGGCCATACCAATGTCTAT δλ g g ij Gaps Score 429.2; DB 22; Length 1367; ä 0; Mismatches 463; Indels Pred. No. 6.6e-98;

60.78;

Best Local Similarity 60.7 Matches 718; Conservative

945 CAGAAGGICIGCCAGCITACATITACCCAAACIGICCATTACIGGAACCIAIGAICIGAA 1004

885 ACAGCACCTGGAAATGAACTCACCCACGATATCATCACCAAGTTCCTGGAAAATGAAGA 851 TCGTCGTAGCGCTTCTCTGCACCTGCCAAAGTTAAGTATCACCGGTACTTACGACTTAAA 1005 GAGCGTCCTGGGTCAACTGGGCATCACTAAGGTCTTCAGCAATGGGGCTGACCTCTCCGG

g δŏ

δ

TGTTACTGAAGAAGCTCCATTAAAATTGAGTAAAGCTGTTCACAAAGCCGTCTTAACTAT

850

790

610

585 CGTGGAGAAGGGTACTCAAGGGAAAATTGTGGATTTGGTCAAGGAGCTTGACAGAGACAC

551 CGTCTTCGCACTAGTTAACTATATTTTTTCAAGGGTAAGTGGGAACGTCCTTTCGAGGT 645 AGTTTTTGCTCTGGTGAATTACATCTTCTTTAAAGGCAAATGGGAGAGACCTTTTGAAGT

TGTTGAGAAAGGCACCCAGGGTAAGATCGTTGACCTAGTTAAAGAATTAGATCGTGATAC

704

730

824

825 GCTAATGAAATACCTGGGCAATGCCACCGCCATCTTCTTCCTACCTGATGAGGGGAAACT 791 TCAACATTTAGAGAATGAGTTGACTCATGACATTATTACTAAATTTTTAGAGAACGAGGA

764

611 TAAAGATACTGAAGAGGAAGATTTTCATGTTGATCAAGTTACTACTGTCAAAGTTCCAAT 705 CAAGGACACCGAGGACGAGGACTTCCACGTGGACCAGGTGACCACGTGAAGGTCCCTAT 671 GATGAAAAGACTGGGTATGTTCAATATTCAACATTGCAAAAAATTAAGTTCTTGGGTCTT

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invention. The specification describes a method for producing mature invention. The specification describes a method for producing mature intention. The specification describes a method for producing mature heterologous protein in monocotyledonous plant cells. The method comprises transforming the cells with a chimeric gene comprising a monocotyledon transcription regulator, inducible either during seed maturation or by adding/removing a small molecule. DNA encoding the heterologous protein, and DNA encoding a signal peptide, with the signal peptide causing secretion of the protein from the cell. Proteins expressed in this manner include mature glycosylated alpha in antirrypain (AMT) with a glycosylation pattern that significantly increases its serum half-life, mature glycosylated antithrombin III (ATIII), mature human serum albumin (HSA) having the native folding capterer as shown by bilitubin-binding characteristics, or mature active subtlinish BDN'. These proteins are useful therapeutically (e.g. AAT for treating emphysema, ATIII and HSA) as blood replacement)
                                                                                                                                                                                                                                                                                                                                                              Protein expression; monocotyledon plant cell; glycosylated antithrombin III; glycosylated alpha 1-antitrypsin; AAT; glycosylated antithrombin III; AATI; phwan serum albumin; HSA; subtilisin BPN'; treatment; emphysema; antithrombotic; blood replacement; ss.
present sequence represents the native coding sequence of mature
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Expressing mature, glycosylated proteins in monocotyledonous cells - from chimeric gene including signal peptide sequence, specifically therapeutic agents and industrial enzymes
                                                                                                                                                                                                                                                                                                                         Native coding sequence of mature alphal-antitrypsin (AAT)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Sequence 1185 BP; 328 A; 324 C; 283 G; 250 T; 0 other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   or as industrial enzymes (BPN' is used in detergents)
                                                          1151 AAGCCCATTGTTTATGGGTAAGGTTGTCAACCCCAACTCAGAA 1192
                                                                               1245 GTCTCCCCTCTTCATGGGAAAAGTGGTGAATCCCACCCAAAA 1286
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Disclosure; Page 29; 53pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (PHYT-) APPLIED PHYTOLOGICS INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             97US-0038170.
97US-0037991.
97US-0038168.
97US-0038169.
                                                                                                                                                                                                  AAV41726 standard; DNA; 1185
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P-PSDB; AAW59839.
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                                                                                                                                                                                                                                                                                 20-NOV-1998
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                                                                                                                                                                                                                                       AAV41726;
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                                                                                                                                                                                 AAV41726
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GATGAAAAGGGTACCGAGGCCGCCGCGCTATGTTCCTGGAAGCTATTCCAATGAGCATT 1091
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       672 ATGAAAAGACTGGGTATGTTCAATATTCAACATTGCAAAAAATTAAGTTCTTGGGTCTTA 731
                                                                            251
                                                                                                                                                                                                                                                                                                                                                                                   372 TCTGAAGGTTTAAAATTGGTTGACAAATTCCTAGAAGACGTCAAGAAACTATATCATAGT 431
                                                             131
                                                                                                                          TTAGCTCATCAAAGTAATTCTACTAACATTTTTTTTTAGTCCTGTTTCTATTGCCACTGCT 191
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71
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              181 GTGGAGAAGGGTACTCAAGGGAAAATTGTGGATTTGGTCAAGGAGCTTGACAGAGACACA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            792 CAACATTTAGAGAATGAGTTGACTCATGACATTATTACTAAATTTTTAGAGAACGAGGAT
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                  CCGACTTTTAATAAAATTACTCCAAATTTAGCCGAATTTGCTTTTTCTTTGTATAGACAA
                                                                                                                                                                                         192 ITCGCCATGTTGAGTTTAGGTACTAAAGCCGATACCCATGACGAGATTTTAGAAGGTTTA
                                                                                                                                                                                                                       181 TTTGCAATGCTCTCCCTGGGGACCAAGGCTGACACTCACGATGAAATCCTGGAGGGCCTG
                                                                                                                                                                                                                                                        AACTITAATITIGACCGAAATCCCAGAAGCCCCAAATTCACGAGGGTTTTCAAGAGTTGTTG
                                                                                                                                                                                                                                                                         312 AGAACTTTGAATCAACCTGATTCTCAATTGCAATTAACTACTGGTAACGGTTTATTTTG
                                                                                                                                                                                                                                                                                                                                                                                                                                                   GAGGCTTTTACCGTTAATTTTGGTGATACTGAGGAAGCTAAAAAGCAAATTAATGATTAT
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GAAGACCCTCAAGGCGACGCCGCTCAAAAAACCGACACCAGTCATCACGACCAAGACCAT
                                                                                                                                                        121 CTGGCACACCAGTCCAACAGCACCAATATCTTCTTCTCCCCAGTGAGCATCGCTACAGCC
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28.1%; Score 429; DB 19; Length 1185; 60.2%; Pred. No. 7.1e-98; 11ve 0; Mismatches 470; Indels 0

Best Local Similarity 60.2 Matches 711; Conservative

Query Match

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192
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    A plasmid consisting of a pCMV4 expression vector including a coding sequence of human alpha-1 antitrypsin may be incorporated into liposomes capable of targeting specific tissue. The plasmid is then capable of expression of the gene extrachromosomally in the cells of the target tissue. Thus, the liposome including the cells of the target tissue. Thus, the liposome including the plasmid can be used in a method for treating a deficiency of the plasmid can be used in a method for treating a deficiency of the specific use of the human alpha-1 antitrypsin is significant as this antiprotease is important in protecting the lungs against to involve a relative deficiency of antiprotease activity.

C Therefore, the delivery of a functioning alpha-1 antiprotease contitions are to the lungs can be therapeutic in many human conditions
1081 CCCCCCGAGGTCAACAACAAAACCCTTTGTCTTAATGATTGAACAAAATACCAAG 1140
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          152 CCAACCTTCAACAAGATCACCCCCAACCTGGCTGAGTTCGCCCTTCAGCCTATACCGCCAG 211
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        132 TTAGCTCATCAAAGTAATTCTACTAACATTTTTTTTAGTCCTGTTTCTATTGCCACTGCT 191
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  92 GAGGATCCCCAGGGAGATGCTGCCCAGAAGACAGATACATCCCCACCATGATCAGGATCAC 151
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          0;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Human alpha-1 anti-trypsin contg. plasmid - for treatment of anti-protease deficiency in emphysema and other lung diseases
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Length 1352;
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Pred. No. 7.4e-98;
0; Mismatches 470; Indels
                                                                                                                                                                                                                                                                     emphysema;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sequence 1352 BP; 349 A; 386 C; 325 G; 292 T; 0 other;
                                            1152 AGCCCATTGTTATGGGTAAGGTTGTCAACCCAACTCAGAA 1192
                                                                      1141 rerecererrearessanastesreareceaeceana 1181
                                                                                                                                                                                                                                                                Plasmid; pCMV4; liposome; antiprotease; lung; adult respiratory distress syndrome; ARDS; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Canonico A, Conary J, Meyrick B;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Disclosure; Fig 6a-6b; 32pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    characterised by injury of the lungs.
                                                                                                                                                AAQ31403 standard; DNA; 1352
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60.2%;
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                                                                                                                                                                                                                                                                                                                                                                                                  92WO-US02465
                                                                                                                                                                                                           (first entry)
                                                                                                                                                                                                                                     Human alpha-1 antitrypsin.
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   WPI; 1992-398857/48.
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                                                                                                                                                                                                                                                                                                                                                                                                                             24-APR-1991;
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212 CTGGCACACCAGTCCAACAACACAATATCTTCTCCCCCAGTGAGCATCGCTACAGCC 271
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                                                                                                         TICGCCATGITGAGITIAGGIACIAAAGCCGAIACCCAIGACGAGAITITAGAAGGITIA 251
                                                                                                                                                                                                                                                          252 AACTTTAATTTGACCGAAATCCCAGAAGCCCAAATTCACGAGGGTTTTCAAGAGTTGTTG 311
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                                                                                                                                                                                                                                                                                                         272 TITGCAATGCTCTCCCTGGGGACCAAGGCTGACACTCACGATGAAATCCTGGAGGGCCTG
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                632 GTTTTTGCTCTGGTGAATTACATCTTTTAAAGGCAAATGGGAGAGACCCTTTGAAGTC
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                                                            Human; cancer; colon; breast; ovary; oesophagus; kidney; thyroid;
stomach; lung; prostate; pancreas; carcinoma; antitumour; cancerous;
cytostatic; gene therapy; antineoplastic; Wilm's tumour; adenocarcinoma;
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Horrigan
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Screening for anti-neoplastic agent involves exposing cells to a
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                                                 Thyroid cancer related gene sequence SEQ ID NO:5848.
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       ABL67511 standard; DNA; 1352 BP
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                                  15-MAY-2002 (first entry)
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Weaver Z;
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The present invention describes a method (M1) for screening for an anti-neoplastic agent. The method involves exposing cells to a chemical agent to be tested for anti-neoplastic activity, determining a change in expression of at least one gene (I) of a signature gene set, where (I) comprises a sequence (S) selected from 8447 sequences (given in ABL61664 to ABL70110), or is at least 95% identical to (S), where a change in cativity and can be used in gene therapy. MI can be used for screening an anti-neoplastic activity. (I) has cytostatic activity and can be used in gene therapy. MI can be used for screening an anti-neoplastic agent, and can be used for producing a product which is the data collected with respect to the anti-neoplastic agent as a result of M1, and the data is sufficient to convey the chemical structure and/or properties of the agent. MI can be used in the treatment of cancer such as colon, breast, stomach, lung, thyroid, cospophageal, ovarian, kidney, prostate or pancreatic cancer, adenocarcinoma, carcinoma, clear cell cancer, infiltrating lobular cancer, squamous cell cancer, infiltrating lobular cancer, squamous cell cancer infiltrating lobular cancer.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          212 CTGGCACACCAGTCCAACAGCAATATCTTCTCTCCCCAGTGAGCATCGCTACAGCC 271
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tested for anti-neoplastic activity, and in expression of a gene of a signature gene set
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             carcinoma, papillary carcinoma and Wilm's tumour.
                                                                                                      Claim 1; SEQ ID 5848; 44pp; English.
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   chemical agent to be determining a change
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